

1/7

FIGURE 1A

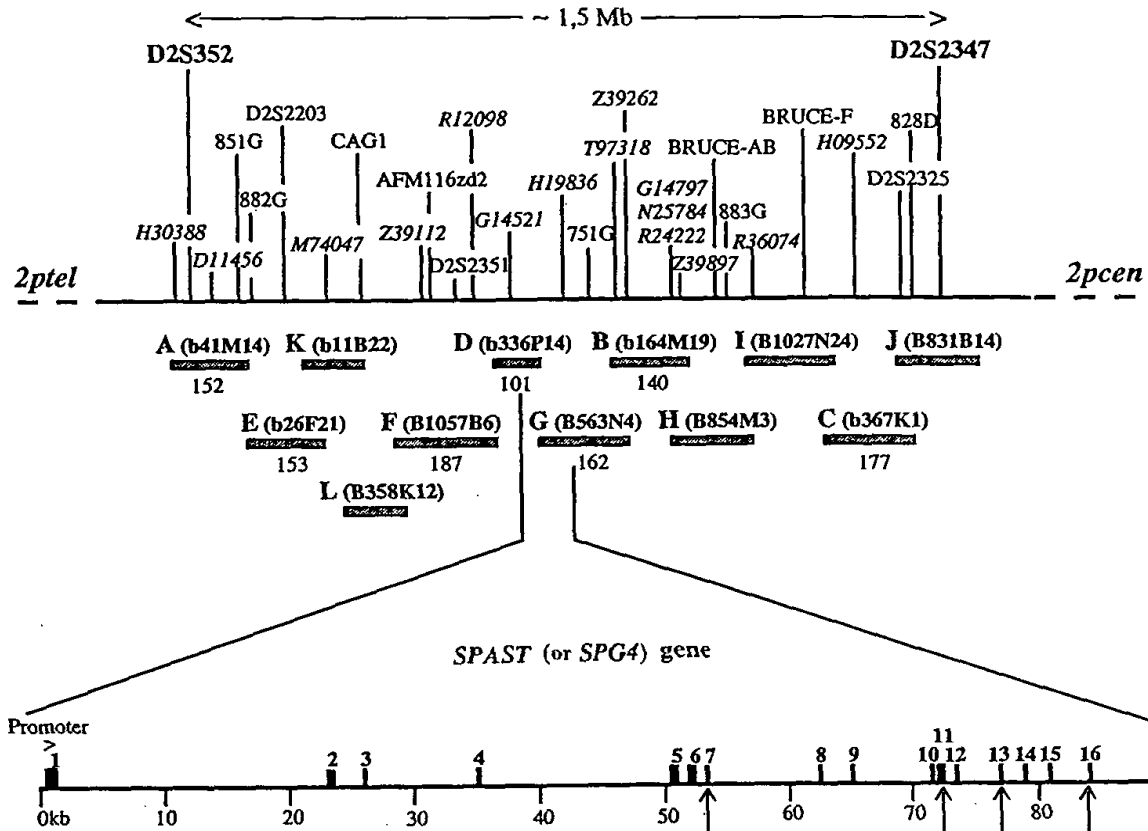
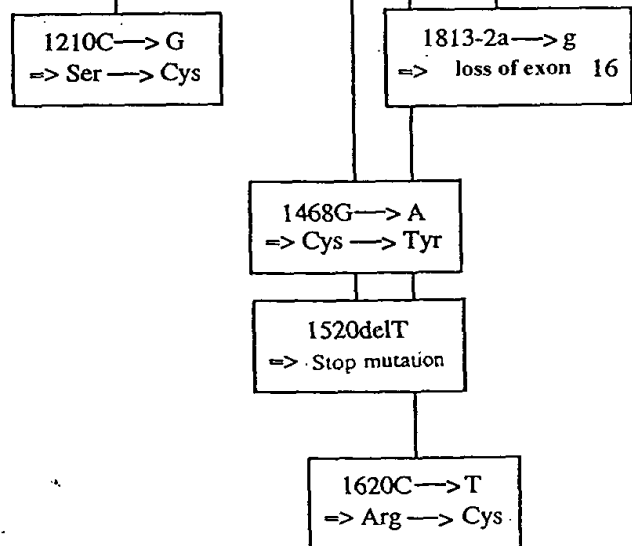


FIGURE 1B

FIGURE 1C



2/7

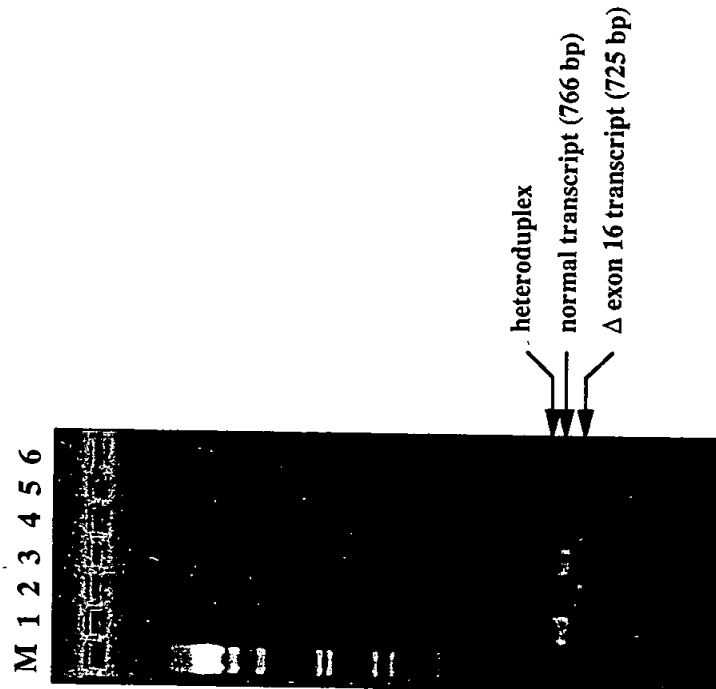
1 GCTCCTGAGACCGGCGGCACACGGGGTCTGTGGCCCCCGCCGTAGCAGTGGCTGCCGCGCTCGCTTGGTTCCCGTGGTCTGCGGGAGGCGGG 95
1 TTATGGCGGCGGCGGCGAGCTGAGAGCTGTAATGAATTTCTCCGGTGGACGAGGGAAGAAAGGCTCCGGCGGCGCCAGCAACCCGGTGCCTCC 190
1 M N S P G G R G K K K G S G G A S N P V P P
CAGGCTCCGCCCCCTTGCCTGGCCCCCGCCCTCCCGCGCGCGGCGGCGCCCTCCGCGCGAGTCGCGCCATAAGCGGAACCTGTACTATTCT 285
23 R P P P C L A P A P A A G P A P P P E S P H K R N L Y Y F S
CCTACCCGCTGTTTGTAGGCTTCGCGCTGCTGCGTTTGGTCCGCTTCCACCTGGGGCTCTCTTTCGTGTGGCTCTGCCAGCGTCTCCCGCGCC 380
55 Y P L F V G F A L L R L V A F H L G L L F V W L C Q R F S R A
CTCATGGCAGCCAGAGGAGCTCCGGGGCGCGCCAGCAGCTGCTCGGCTCGGCGCGCGCGCGCGGTGCCGGGCGGCGAGGCGCGAGCGCGTCCG 475
86 L M A A K R S S G A A P A P A S A S A P A P V P G G E A E R V R
AGTCTTCCACAACAGGCGCTTCGAGTACATCTCCATTGCGCTGCGCATCGATGAGGATGAGAAAGTGGACAGAAGGAGCAAGCTGTGGAATGGT 570
118 V F H K Q A F E Y I S I A L R I D E D E K A 2 G Q K E Q A V E W Y
ATAAGAAAGGTATTGAAGAACTGGAAAAAGGAATAGCTGTTATAGTITACAGGACAAGTGAACAGTGTGAAGAGCTAGACGCTTCAAGCTAAA 665
150 K K G I E E L E K G I A V I V T G Q G 3 E Q C E R A R R L Q A K
ATGATGACTAATTTGGTTATGGCCAAGGACCGCTTACAACCTTCTAGTGAAGATGCAACAGTTTTGCCATTTTCCAAGTCAAAAACGGACGTCTA 760
181 M M T N L V M A K D R L Q L L E K M Q P V L P F S K S Q T D V Y
TAATGACAGTACTAATTTGGCATGCCGCAATGGACATCTCCAGTCAAGTGGAGCTGTTCCAAAAAGAAAAGACCCCTTAACACACACTAGTA 855
213 N D S T N L A C R N H A P A S E S A S A P A P V P K R K D P L T H T S N
ATTCAGTGCCTCGTTCAAAAACAGTTATGAAAACCTGGATCTGCAGGCGCTTTCAGGCCACCATAGAGCACCTAGTTACAGTGGTTTATCCATGGTT 950
245 S L P R S K T V M K T G S A G L S G H H R A P S Y S G L S M V
TCTGGAGTGAACAGGAGTCTGCTCTGCTCTCTAAGTGTACTCCGAAAACAAATAGGACAAAATAACCTTACCCCTTACAACCTGC 1045
276 S G V K Q G S G P A P T T T H K G 6 T P K T N R T N K P S T A C C P T T A
TACTCGTAAGAAAAAGACTGAAGAATTTTAGGAATGTGGACAGCAACCTTGCTAACCTTATAATGAATGAAATGTGGACAATGGAAACAGCTG 1140
308 T R K K K D L K N F R N V D S N L A N L I M N E I V D N G 7 T A V
TTAAATTTGATGATATAGCTGGTCAAGACTTGGCAAAACAGCATTGAAGAAATTTGTTATTTCTTCTCTGAGGCGCTGAGTTGTTCAAGGG 1235
340 K F D D I A G Q D L A K Q A L Q E I V I L P S L R P E L 8 F T G
CTTAGAGCTCCTGCCAGAGGCTGTACTCTTTGGTCCACTGGGAATGGGAAGACAATGCTGCTAAAGCAGTAGCTGCGAATCGAATGCCAAC 1330
371 L R A P A R G L L L F G P P G N G K T M L A 9 K A V A A E S N A T
CTTCTTTAATATAAGTGTGCAAGTTTAACTTCAAAATATCTGGGAGAGGAGAGAAATTTGGTGGAGGCTCTTTTGGCTGGGCTCGAGAACTTC 1425
403 F F N I S A A S L T S K Y 10 G E G E K L V R A L F A V A R E L Q
AACCTTCTATAATTTTATAGTGAAGTTGATAGCCTTTTGTGTGAAAGAGAGAGGGGAGCAGATGCTAGTAGACGCTTAAAACTGAATTT 1520
435 P S I I F I D I E V D S L L C E R R E G E H D A S R R L K T E F
CTAATAGAAATTTGATGGTGTACAGTCTGCTGGAGATGACAGAGTACTTGAATGGGTGCAACTAATAGGCCACAAGAGCTTGTATGAGGCTGTCT 1615
466 L I E F D G 12 Q S A G D D R V L V M G A T N R P Q E L D E A V L
CAGGCTTTTCATCAACCGGCTATATGTGTCTTTACCAAAATGAGGAGCAAGACTACTTTTGTCTAAAAATCTGTTATGTAAACAAGGAAGTCCAT 1710
498 R 13 R F I K R V Y V S L P N E E 14 R L L L L K N L L C K Q G S P L
TGACCCAAAAGAACTAGCACAACTTGCTAGTATGACTGATGGATCTCAGGAAGTACCTAACAGCTTTGGCAAAAGATGCAGCACTGGGTCT 1805
530 T Q K E L A Q L A R M 15 T D G Y S G S D L T A L A K D A A L G P
ATCCGAGTAACTAAAACAGAAACAGGTGAAGAATATGTCTGCCAGTGAATGAGAAATATTCGATTATCTGACTTCACTGAATCCTTGAAAAAAT 1900
561 I R E 16 L K P E Q V K N M S A S E 17 R N I R L S D F T E S L K K I
AAAACGAGCGTCAAGCCTCAAACTTTAGAAGCGTACATACGTTGGAACAAGGACTTTGGAGATACCCTGTTTAAGGAAATACCTTTGTAAACC 1995
593 K R S V S P Q T L E A Y I R W N K D F G D T T V *
TGCAGAACATTTTACTTAAAGAGGAAACACAAGATCTTCAATGAACGTCATCGGCTACAGAAACAGCCTAAGTTTACAGGACTTTTTAGAGTCT 2090
TACATATTTGTGCACCAAACTTGAAGATGAACCAAGAAACAGACTTAAACAAAATATACAATGCAAAATGTAATTTTGTGTGTTAAGGCCCTTGC 2185
CTTGATGGTCACAGTTATCCCAATGGACACTAAGTTAGAGCACAAACAAACCTGATTCTGGTCTTCTTTACCAATATAATCATAATGTAATAAT 2280
AATTTGTATATTGTTTGCAGATGAAAGTATTCCAGGAACAGTGAATGGTAGAAGACACAAGAACATTGTTTGTGTTCTCTGATGTTTTTTC 2375
TTAAAAATAGTAATTTCTCTACTTTTCTTTCTACTGTTGCTTAACTACAGGTGATGGAATGCCAAACACTCTTAAGTTTATTTTCTTTTTC 2470
GTTTTATAAATCAGTGTGCCAAATGAAACTTTTCTTAAGTAACTGTAATAGGAAAAAGTTTATTTTGAGAGTTCTTCTTCATAAATCTACA 2565
GACATTAACAATTTGTTGTGTTCTTTTACCTTTTATTTTCTATTACCTTGCTACCAACAGTITAGATAGCAATATAATAGCAAAAAAGCAAA 2660
TATGTTAAAAATAGAGAAGGTTTGAAGGTTTGTAGTTACTCTGCTATATAACATGTAGATCAGTCTTCATGTGACCTGCAGTATTTTTTTCTAAT 2755
GTATTTGTGCAAAATCTGTTGTAGACTGTTAACTTCTCTGCTGGAATTTATTTCTGCAAGAATTATTTCTGATATTTAAGAGAGCCAATTTTA 2850
ACTGCTGTGAAAAATGTTTCCAGTGAAGAGAGGAAATACTAGGAACAAAGACATTTCTAATTTATTGCTTATTACTTTCTTAATTTTACAGGA 2945
TAATTATAAGCAAGTGAACCTACCTTTTATCTTAATAATTATTAATCCCTTCAATGAACTTTAAAAAACTGAATTTTATACATGGCAT 3040
ACATTTTCTAGTTCCTTCTGCTTCTTTATTAACCTAAAAGTTCTAGTTCTAGTCTGTTGATCTGCCTTTTGTCTCCCAAAATGTACAGTAAT 3135
TCCATTTGTTGTATAAATATGCTTGGATTTTCAATATAAATATGTCATTGTAGGAGTAGAGACTCATATCATGGCCTTTTAAATATTGTAATA 3230
AAGGCAAAATAGATATTTGCCCTTAGTTACTGG 3263

FIGURE 2

09/830902

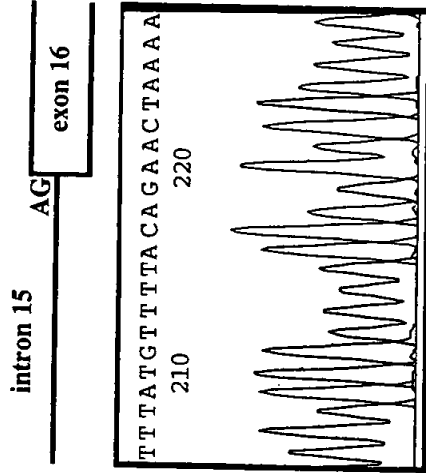
3/7

A



B

Normal acceptor splice site



Mutated acceptor splice site

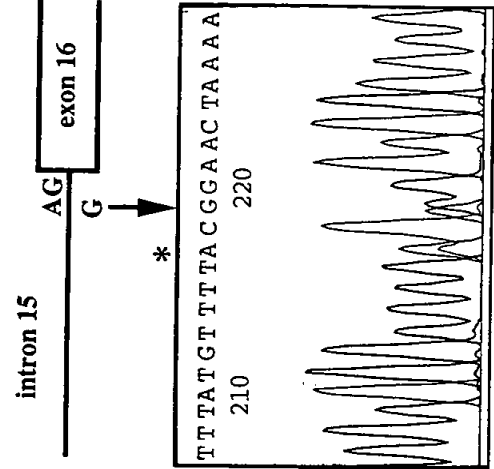


FIGURE 3

FIGURE 4A

[illegible]

FIGURE 4B

[illegible]

6/7

Human: 1 [...] 459
Mouse: 1 AGGCCGAGAGCGTCCGCGTCTTCCACAAGCAGGCCTTCGAGTACATCTCCATTGCCCTGC 60
Human: 460 AGGCCGAGCGCGTCCGAGTCTTCCACAAACAGGCCTTCGAGTACATCTCCATTGCCCTGC 519
Mouse: 61 GCATCGACGAGGAAGAGAAAGCAGGACAGAAGGAACAAGCTGTGAATGGTATAAGAAAG 120
Human: 520 GCATCGATGAGGATGAGAAAGCAGGACAGAAGGAGCAAGCTGTGAATGGTATAAGAAAG 579
Mouse: 121 GTATCGAAGAACTGGAAAAAGGAATCGCTGTTATAGTTACGGGCCAAGGTGAACAGTATG 180
Human: 580 GTATTGAAGAACTGGAAAAAGGAATAGCTGTTATAGTTACAGGACAAGGTGAACAGTGTG 639
Mouse: 181 AAAGAGCTAGACGCTCTCAAGCCAAAATGATGACTAATTAGTTATGGCCAAGGACCGCTT 240
Human: 640 AAAGAGCTAGACGCTCTCAAGCTAAAATGATGACTAATTGGTTATGGCCAAGGACCGCT 699
Mouse: 241 TACAACCTCTAGAGAAGCTGCAACCAAGTTTGCAATTTTCCAAGTCACAGACGGACGCTCT 300
Human: 700 TACAACCTCTAGAGAAGATGCAACCAAGTTTGCCATTTTCCAAGTCACAAACGGACGCTCT 759
Mouse: 301 ATAACGAGAGTACTAACCTGACATGCCGCAATGGACATCTCCAGTCAGAAAGTGGAGCAG 360
Human: 760 ATAATGACAGTACTAACTTGGCATGCCGCAATGGACATCTCCAGTCAGAAAGTGGAGCTG 819
Mouse: 361 TTCCGAAGAGGAAGACCCCTTAACACATGCTAGTAATTCATTGCTCGATCAAAACATG 420
Human: 820 TTCCAAAAAGAAAGACCCCTTAACACACACTAGTAATTCAGTGCCTCGTTCAAAACAG 879
Mouse: 421 TCCTGAAAAGTGGCTCCGCGAGGCTCTCCGGTCACACAGGCGCTAGTTGCAGTGGTT 480
Human: 880 TTATGAAAAGTGGATCTGCAGGCTTTCCAGGCCACCATAGAGCACTAGTTACAGTGGTT 939
Mouse: 481 TGTCCATGTTTCTGGAGCAAGACCGGGACCTGGTCTGCAGCTACCAACATTAAGGTA 540
Human: 940 TATCCATGTTTCTGGAGTGAACAGGGATCTGGTCTGCTCTACCACTCATAAGGTA 999
Mouse: 541 CTCCAAAACCAATAGAAACCAACCACTTCTACTCCCACTGCAAGTTCGGAAAAAGA 600
Human: 1000 CTCCGAAAACCAATAGGACAAATAAACCTTCTACCCCTACAACTGCTACTCGTAAGAAA 1059
Mouse: 601 AAGACTTGAATAATTTAGGAATGTGGACAGCAATCTTGCTAACCTTATAATGAATGAAA 660
Human: 1060 AAGACTTGAATAATTTAGGAATGTGGACAGCACTTGCTAACCTTATAATGAATGAAA 1119
Mouse: 661 TTGTTGACAAATGGGACAGCTGTTAAGTTGATGACATAGCCGGGAGGAGCTGGCAAGC 720
Human: 1120 TTGTTGACAAATGGAAACAGCTGTTAATTTGATGATATAGCTGGTCAAGACTTGGCAAAAC 1179
Mouse: 721 AAGCGCTGAGGAGATTTGTCATCCTTCTCTCTGCGGCTGAGTTGTTACAGGGCTCA 780
Human: 1180 AAGCATTGCAAGAAATTTGTTATCTCTCTCTGAGGCTGAGTTGTTACAGGGCTTA 1239
Mouse: 781 GAGCTCCTGCTAGAGGCTTGTACTCTTCCGGTCCGCCAGGAAACGAAAAACAATGCTGG 840
Human: 1240 GAGCTCCTGCCAGAGGGCTGTACTCTTGGTCCACCTGGGAATGGGAAGACAATGCTGG 1299
Mouse: 841 CTAAGCAGTAGCTGCAGAGTCTAATGCGACCTTTTCAACATAAGTCTGCCAGTTAA 900
Human: 1300 CTAAGCAGTAGCTGCAGAAATCGAATGCAACCTTCTTAATATAAGTCTGCAAGTTAA 1359
Mouse: 901 CTTCAAAATATGTGGGAGAGGAGAGAAATGGTGAGAGCTCTCTTTGCTGTGGCTCGAG 960
Human: 1360 CTTCAAAATACGTGGGAGAGGAGAGAAATGGTGAGGGCTCTTTTGTGTGGCTCGAG 1419
Mouse: 961 AACTTCAACCATCTATAATTTTATAGATGAAGTTGACAGCTTTTGTGTGAGAGACGGG 1020
Human: 1420 AACTTCAACCTTCTATAATTTTATAGATGAAGTTGATAGCCTTTTGTGTGAAGAAGAG 1479
Mouse: 1021 AAGGGGAGCAGCAGCTAGCAGACGGCTAAAGACGGAATTTTAATAGAATTTGACGGGG 1080
Human: 1480 AAGGGGAGCAGCAGTCTAGTAGACGCTAAAACTGAATTTCTAATAGAATTTGATGGTG 1539
Mouse: 1081 TGCAATCTGCTGGAGATGACAGAGTACTGTAATGGGTGCAACTAACAGGCCCAAGAGC 1140
Human: 1540 TACAGTCTGCTGGAGATGACAGAGTACTGTAATGGGTGCAACTAATAGGCCACAAGAGC 1599
Mouse: 1141 TTGATGAAGCTGTTCTCAGGCGTTTCATTAAACGGGTATATGTGCTTACCAATGAGG 1200
Human: 1600 TTGATGAGGCTGTTCTCAGGCGTTTCATCAACGGGTATATGTGCTTTACCAATGAGG 1659
Mouse: 1201 AGACAAGACTCCTTCTGCTTAAAAACCTGTTGTGTAACAAGGAAGTCCACTGACCCAAA 1260
Human: 1660 AGACAAGACTACTTTTGCTTAAAAATCTGTTATGTAAACAAGGAAGTCCATTGACCCAAA 1719
Mouse: 1261 AAGAACTCGCACAGCTTGTCTAGAAATGACCGATGGTACTCTGGAAGTGTCTGACCGCTT 1320
Human: 1720 AAGAACTAGCACAACCTTGCTAGAAATGACTGATGGTACTCAGGAAGTGACCTAACAGCTT 1779
Mouse: 1321 TGGCCAAGGATGCAGCCCTGGGTCTATCCGAGAAGTGAAGCCAGAGCAGGTGAAGAATA 1380
Human: 1780 TGGCAAAAGATGCAGCACTGGGTCTATCCGAGAAGTGAAGCCAGAGCAGGTGAAGAATA 1839
Mouse: 1381 TGTCTGCCAGTGAGATGAGAAATATTCGATTATCTGACTTCACAGAATCCTTAAAAAGA 1440
Human: 1840 TGTCTGCCAGTGAGATGAGAAATATTCGATTATCTGACTTCACTGAATCCTTGAAGAAAA 1899
Mouse: 1441 TAAACGCAAGTGTGAGTCTCAGACCTTAGAAGCATACATACGCTGGAACAAGGATTTG 1500
Human: 1900 TAAACGCAAGCTGACGCCCTCAAACTTTAGAAGCGTACATACGTTGGAACAAGGACTTTG 1959
Mouse: 1501 GAGACACCACTGTTTAAAGGAAT 1523
Human: 1960 GAGATACCACTGTTTAAAGGAAT 1982
Human: 1983 [...] 3263
Mouse: 1524 GGATGCCTCTGTGAGCCCATAGAACATCGCACTTCACAGGAACAAGAGCTTTGGCTACA 1583
1584 GGAACCCAGACTTCGTTTACAGGACGTTTATAGATTTTCATTTTGTGCACCAAACTTGA 1643
1644 AGAGGAACAAGAAGACAGACCTAAATAAATATGCAATATGAATGG 1689

FIGURE 5

106101-2060E860

FIGURE 6B

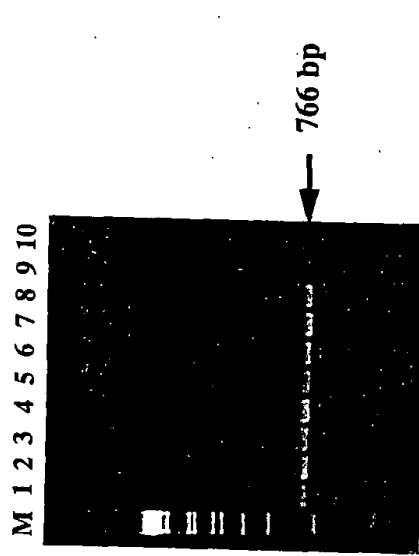


FIGURE 6C

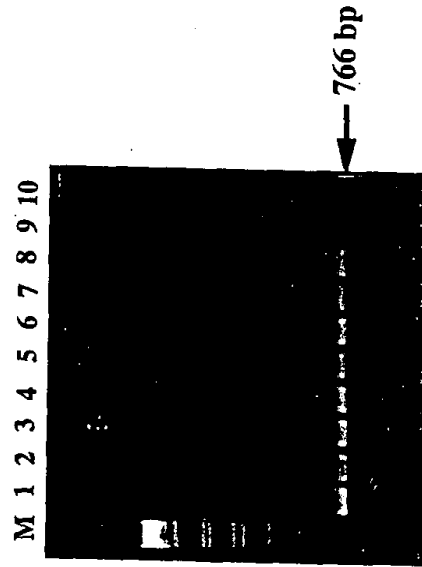


FIGURE 6A

